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- 1. A method for generating a secondary library of scaffold protein variants comprising:
  - a) providing a primary library comprising a filtered set of scaffold protein primary variant sequences;
  - b) generating a list of primary variant positions in said primary library;
  - c) combining a plurality of said primary variant positions to generate a secondary library of secondary sequences.
- 2. A method for generating a secondary library of scaffold protein variants comprising:
  - a) providing a primary library comprising a filtered set of scaffold protein primary variant sequences;
  - b) generating a probability distribution of amino acid residues in a plurality of variant positions;
  - c) combining a plurality of said amino acid residues to generate a secondary library of secondary sequences.
- 3. A method according to claim 1 further comprising synthesizing a plurality of said secondary sequences.
- 4. A method according to claim 2 wherein said synthesizing is done by multiple PCR with pooled oligonucleotides.
- 5. A method according to claim 4 wherein said pooled oligonucleotides are added in equimolar amounts.
- 6. A method according to claim wherein said pooled oligonucleotides are added in amounts that correspond to the frequency of the mutation.
- 7. A composition comprising a plurality of secondary variant proteins comprising a subset of said secondary library.
- 8. A composition comprising a plurality of nucleic acids encoding a plurality of secondary variant proteins comprising a subset of said secondary library.
- 9. A method for generating a secondary library of scaffold protein variants comprising:
  - a) providing a first library fiftered set of scaffold protein primary variants;
  - b) generating a probability distribution of amino acid residues in a plurality of variant positions;
  - c) synthesizing a plurality of scaffold protein secondary variants comprising a plurality of said amino acid residues to forma secondary library;

wherein at least one of said secondary variants is different from said primary variants.

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